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Gonorrhea Curriculum

Gonorrhea

Neisseria gonorrhoeae

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Gonorrhea Curriculum

Learning Objectives

Upon completion of this content, the learner will be able to

- Describe the epidemiology of gonorrhea in the U.S.
- Describe the pathogenesis of *Neisseria gonorrhoeae*.
- Discuss the clinical manifestations of gonorrhea.
- Identify common methods used in the diagnosis of gonorrhea.
- List CDC-recommended treatment regimens for gonorrhea.
- Summarize appropriate prevention counseling messages for patients with gonorrhea.
- Describe public health measures for the prevention of gonorrhea.

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Gonorrhea Curriculum

Lessons

- I. Epidemiology: Disease in the U.S.
- II. Pathogenesis
- III. Clinical manifestations
- IV. Diagnosis
- V. Patient management
- VI. Prevention

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Lesson I: Epidemiology: Disease in the U.S.

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Gonorrhea Curriculum Epidemiology

Incidence and Prevalence

- Significant public health problem in U.S.
- Number of reported cases underestimates incidence
- Incidence remains high in some groups defined by geography, age, race/ethnicity, or sexual risk behavior
- Increasing proportion of gonococcal infections caused by resistant organisms

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Gonorrhea—Rates, United States, 1941–2011

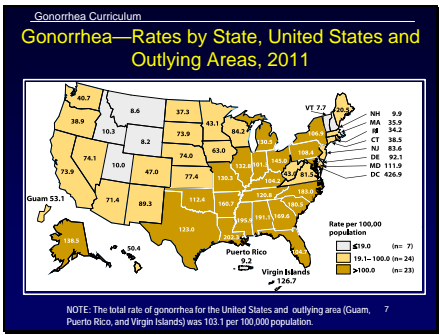
Rate (per 100,000 population)

Year

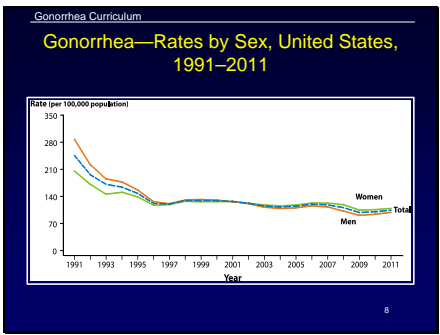
Year	Rate (per 100,000 population)
1941	150
1945	250
1950	150
1955	150
1960	150
1965	200
1970	350
1975	450
1980	450
1985	350
1990	200
1995	150
2000	150
2005	150
2010	150

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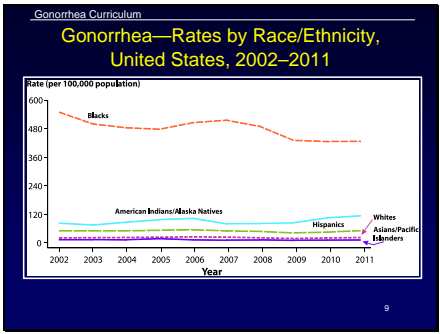
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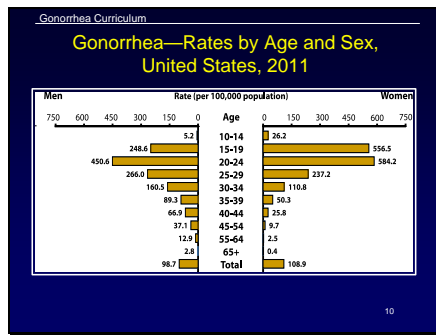
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Epidemiology

Risk Factors

- Multiple or new sex partners or inconsistent condom use
- Urban residence in areas with disease prevalence
- Adolescent, females particularly
- Lower socio-economic status
- Use of drugs
- Exchange of sex for drugs or money
- African American

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Epidemiology

Transmission

- Efficiently transmitted by
 - Male to female via semen
 - Vagina to male urethra
 - Rectal intercourse
 - Fellatio (pharyngeal infection)
 - Perinatal transmission (mother to infant)
- Gonorrhea associated with increased transmission of and susceptibility to HIV infection

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Lesson II: Pathogenesis

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Gonorrhea Curriculum Pathogenesis

Microbiology/Pathology

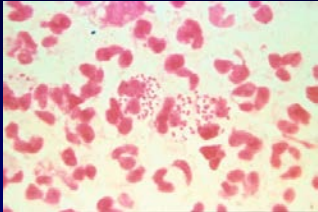
- Etiologic agent: *Neisseria gonorrhoeae*
- Gram-negative intracellular diplococcus
- Infects mucus-secreting epithelial cells
- Evades host response through alteration of surface structures

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Gonorrhea Curriculum Pathogenesis

Gonorrhea: Gram Stain of Urethral Discharge



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Source: CDC/NCHSTP/Division of STD Prevention, STD Clinical Slides

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Lesson III: Clinical Manifestations

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Gonorrhea Curriculum Clinical Manifestations

Genital Infection in Men

- Urethritis – Inflammation of urethra
- Epididymitis – Inflammation of the epididymis

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Gonorrhea Curriculum Clinical Manifestations

Male Urethritis

- Symptoms
 - Typically purulent or mucopurulent urethral discharge
 - Often accompanied by dysuria
 - Discharge may be clear or cloudy
- Asymptomatic in a minority of cases
- Incubation period: usually 1-14 days for symptomatic disease, but may be longer

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Gonorrhea Curriculum Clinical Manifestations

Gonococcal Urethritis: Purulent Discharge



Source: Seattle STD/HIV Prevention Training Center at the University of Washington;
Connie Celum and Walter Stamm

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Gonorrhea Curriculum Clinical Manifestations

Epididymitis

- Symptoms: unilateral testicular pain and swelling
- Infrequent, but most common local complication in males
- Usually associated with overt or subclinical urethritis

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Gonorrhea Curriculum Clinical Manifestations

Swollen or Tender Testicle (Epididymitis)



Source: Seattle STD/HIV Prevention Training Center at the University of Washington

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Clinical Manifestations

Genital Infection in Women

- Most infections are asymptomatic
- Cervicitis – inflammation of the cervix
- Urethritis – inflammation of the urethra

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Clinical Manifestations

Cervicitis

- Non-specific symptoms: abnormal vaginal discharge, intermenstrual bleeding, dysuria, lower abdominal pain, or dyspareunia
- Clinical findings: mucopurulent or purulent cervical discharge, easily induced cervical bleeding
- At least 50% of women with clinical cervicitis have no symptoms
- Incubation period unclear, but symptoms may occur within 10 days of infection

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Clinical Manifestations

Gonococcal Cervicitis



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Source: CDC/NCHSTP/Division of STD Prevention, STD Clinical Slides

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Clinical Manifestations

Urethritis

- Symptoms: dysuria, however, most women are asymptomatic
- 70%–90% of women with cervical gonococcal infection may have urethral infection

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Clinical Manifestations

Complications in Women

- Accessory gland infection
 - Bartholin's glands
 - Skene's glands
- Pelvic Inflammatory Disease (PID)
 - May be asymptomatic
 - May present with lower abdominal pain, discharge, dyspareunia, irregular menstrual bleeding and fever
- Fitz-Hugh-Curtis Syndrome
 - Perihepatitis

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Clinical Manifestations

Bartholin's Abscess



Source: CDC/NCHSTP/Division of STD Prevention, STD Clinical Slides

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Clinical Manifestations

Syndromes in Men and Women

- **Anorectal infection**
 - Usually acquired by anal intercourse
 - Usually asymptomatic
 - Symptoms: anal irritation, painful defecation, constipation, scant rectal bleeding, painless mucopurulent discharge, tenesmus, and anal pruritus
 - Evaluate utilizing an anoscopic examination
 - Signs: mucosa may appear normal, or purulent discharge, erythema, or easily induced bleeding may be observed with anoscopic exam
- **Pharyngeal infection**
 - May be sole site of infection if oral-genital contact is the only exposure
 - Most often asymptomatic, but symptoms, if present, may include pharyngitis, tonsillitis, fever, and cervical adenitis

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Clinical Manifestations

Syndromes in Men and Women (continued)

- **Conjunctivitis**
 - Usually a result of autoinoculation in adults
 - Symptoms/signs: eye irritation with purulent conjunctival exudate
- **Disseminated gonococcal infection (DGI)**
 - Systemic gonococcal infection
 - Occurs infrequently. More common in women than in men
 - Associated with a gonococcal strain that produces bacteremia without associated urogenital symptoms
 - Clinical manifestations: skin lesions, arthralgias, tenosynovitis, arthritis, hepatitis, myocarditis, endocarditis, and meningitis


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Clinical Manifestations

Gonococcal Ophthalmia



Source: CDC/NCHSTP/Division of STD Prevention, STD Clinical Slides

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Gonorrhea Curriculum Clinical Manifestations

Gonococcal Infection in Children

- Perinatal: infections of the conjunctiva, pharynx, respiratory tract or anal canal
- Older children (>1 year): considered possible evidence of sexual abuse
- Vulvovaginitis, not cervicitis, in prepubescent girls
- Anorectum or pharynx more commonly infected in boys than urethra
- Because of legal implications, culture remains the preferred method of diagnosis

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Lesson IV: Diagnosis

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Diagnosis

Diagnostic Methods

- Culture tests
 - Advantages: low cost, suitable for a variety of specimen sites, antimicrobial susceptibility can be performed
 - Anatomic sites to test: in response to exposure history in persons at significant risk of gonococcal infection, complaints, or clinical findings
 - In men: urethra in all men; pharynx and rectum, depending on exposure history or symptoms
 - In women: cervix should be tested; pharynx and rectum depending on symptoms and exposure history; vagina may be tested if cervix is absent; Bartholin's or Skene's glands may be cultured if overt exudate is expressed

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Diagnosis

Diagnostic Methods (continued)

- Non-culture tests
 - Amplified tests (NAATs)
 - Polymerase chain reaction (PCR) (Roche Amplicor)
 - Transcription-mediated amplification (TMA) (Gen-Probe Aptima)
 - Strand displacement amplification (SDA) (Becton-Dickinson BD ProbeTec ET)
 - Non-amplified tests
 - DNA probe (Gen-Probe PACE 2, Digene Hybrid Capture II)
 - Gram-stained smear

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Diagnosis

Clinical Considerations

- In cases of suspected sexual abuse
 - Adults
 - NAATs are preferred for diagnostic evaluation of sexual assault regardless of penetration
 - Children
 - Culture remains the preferred method for urethral specimens or urine from boys and for extragenital specimens for all children
 - NAATs can be used as an alternative to culture with vaginal specimens or urine from girls

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Lesson V: Patient Management

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Gonorrhea CurriculumManagement

Antimicrobial Susceptibility of *N. gonorrhoeae*

- Fluoroquinolone resistance is widely disseminated throughout the U.S. and the world
- Approximately 25% of isolates are resistance to penicillin or tetracycline or both
- In 2011, 0.3% of isolates showed decreased susceptibility to azithromycin, down from 0.5% in 2010.
- Sporadic cases of decreased susceptibility to ceftriaxone and cefixime have been reported recently

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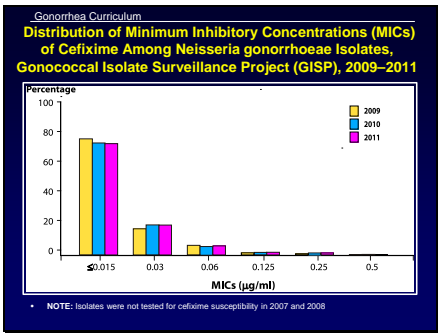
Percentage of *Neisseria gonorrhoeae* Isolates that are Ciprofloxacin-Resistant by Sex of Sex Partner, Gonococcal Isolate Surveillance Project (GISP), 1995–2011

Year	MSM (%)	MSW (%)
1995	0	0
1996	0	0
1997	0	0
1998	0	0
1999	0	0
2000	0	0
2001	0	0
2002	5	0
2003	15	0
2004	25	0
2005	30	5
2006	40	10
2007	35	10
2008	35	10
2009	20	5
2010	25	10
2011	25	10

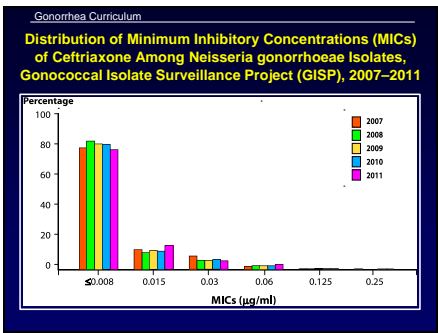
*MSM—men who have sex with men; MSW—men who have sex with women only.

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Gonorrhea Curriculum Management

Treatment for Uncomplicated Gonococcal Infections of the Cervix, Urethra, and Rectum

Recommended

Ceftriaxone	250 mg	IM	Once
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PLUS

Azithromycin	1 g	Orally	Once
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OR

Doxycycline	100 mg	Orally	Twice a day for 7 days
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Quinolones are no longer recommended in the United States for the treatment of gonorrhea and associated conditions, such as PID.

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Gonorrhea Curriculum

Management

Treatment for Uncomplicated Gonococcal Infections of the Cervix, Urethra, and Rectum

Alternative 1: *If Ceftriaxone is not available*

Cefixime	400 mg	Orally	Once
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PLUS

Azithromycin	1 g	Orally	Once
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OR

Doxycycline	100 mg	Orally	Twice a day for 7 days
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PLUS

Test of cure in 1 week			
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Management

Treatment for Uncomplicated Gonococcal Infections of the Cervix, Urethra, and Rectum

Alternative 2: *If patient is cephalosporin-allergic*

Azithromycin	2 g	Orally	Once
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PLUS

Test of cure in 1 week			
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Management

Treatment for Uncomplicated Gonococcal Infections of the Pharynx

Ceftriaxone	250 mg	IM	Once
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PLUS

Azithromycin	1 g	Orally	Once	OR
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Doxycycline	100 mg	Orally	Twice a day for 7 days	
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Gonorrhea CurriculumManagement

Special Considerations:
Pregnancy

- Treat with recommended cephalosporin-based combination therapy
- If cephalosporin is not tolerated, treat with azithromycin 2 g orally. A test of cure should be performed 1 week after treatment
- Pregnant women should not be treated with quinolones or tetracyclines. Spectinomycin is not commercially available

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Gonorrhea CurriculumManagement

Penicillin-Allergic

- Azithromycin 2 g orally
 - Plus test of cure in 1 week
- Desensitization is impractical in most settings

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Follow-Up

- A test of cure is not recommended if recommended regimen is administered
- A test of cure is recommended if an alternative regimen is administered
- If symptoms persist, perform culture for *N. gonorrhoeae*
 - Any gonococci isolated should be tested for antimicrobial susceptibility at site of exposure
- Repeat testing in 3 months

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Lesson VI: Prevention

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Gonorrhea Curriculum Prevention

Screening

- **Pregnancy:**
 - A test for *N. gonorrhoeae* should be performed at the 1st prenatal visit for women at risk or those living in an area in which the prevalence of *N. gonorrhoeae* is high
 - Repeat test in the 3rd trimester for those at continued risk
- **U.S. Preventive Service Task Force** recommends screening all sexually active women for gonorrhea infection if they are at increased risk of infection
- **Sexually active men who have sex with men:** CDC recommends screening at least annually at all anatomic sites of exposure

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Gonorrhea Curriculum Prevention

Partner Management

- Evaluate and treat all sex partners for *N. gonorrhoeae* and *C. trachomatis* infections, if contact was within 60 days of symptoms or diagnosis
- If a patient's last sexual intercourse was >60 days before onset of symptoms or diagnosis, the patient's most recent sex partner should be treated
- Avoid sexual intercourse until therapy is completed and both partners no longer have symptoms

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Prevention

Reporting

- Laws and regulations in all states require that persons diagnosed with gonorrhea are reported to public health authorities by clinicians, labs, or both

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Gonorrhea Curriculum

Prevention

Patient Counseling/Education

- Nature of disease
 - Usually symptomatic in males and asymptomatic in females
 - Untreated infections can result in PID, infertility, and ectopic pregnancy in women and epididymitis in men
- Transmission issues
 - Efficiently transmitted
- Risk reduction
 - Utilize prevention strategies

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Case Study

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Gonorrhea CurriculumCase Study



History: Robert Forbes

- 33-year-old male who presents to his doctor reporting a purulent urethral discharge and dysuria for 3 days.
- Lives in Dallas with history of travel to Las Vegas 3 weeks ago.
- New female sex partner (Laura) for 2 months. They have unprotected vaginal intercourse 4 times/week, the last time being 2 days ago. No oral or rectal sex.
- Also had a one-time sexual encounter with a woman he met in Las Vegas 3 weeks ago (Monica). They had oral and vaginal sex. No condoms used.
- No history of urethral discharge or STDs, no sore throat or rectal discomfort. Negative HIV test 1 year ago.

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Gonorrhea CurriculumCase Study

Physical Exam

- Vital signs: blood pressure 98/72, pulse 68, respiration 14, temperature 37.2° C
- Cooperative, good historian
- Chest, heart, musculoskeletal, and abdominal exams within normal limits
- No flank pain on percussion, normal rectal exam, no sores or rashes
- The genital exam reveals a reddened urethral meatus with a purulent discharge, without lesions or lymphadenopathy

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Gonorrhea CurriculumCase Study

Questions

1. What should be included in the differential diagnosis?
2. Which laboratory tests are appropriate to order or perform?
3. What is the appropriate treatment regimen?

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Gonorrhea CurriculumCase Study

Laboratory Results

Results of laboratory tests:

- Urethral and pharyngeal culture: showed growth of a Gram-negative diplococcus that was oxidase-positive. Biochemical and FA conjugate testing confirmed this isolate to be *N. gonorrhoeae*.
- NAAT for chlamydia: negative
- RPR: nonreactive
- HIV antibody test: negative

4) What is the diagnosis, based on all available information?


5) Who is responsible for reporting this case to the local health department?

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Gonorrhea CurriculumCase Study

Partner Management



Robert's sex partners within the past 3 months:

- Laura: Last exposure - Unprotected vaginal sex 2 days ago
- Monica: Last exposure - Unprotected oral (Monica performed fellatio) and vaginal sex 3 weeks ago while he was in Las Vegas
- Jerilyn: Last exposure - Unprotected vaginal sex 3 months ago



6) Laura was examined and her lab results came back negative for gonorrhea and chlamydia. How should Laura be managed?

7) What tests should Jerilyn and Monica have?

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Gonorrhea CurriculumCase Study

Follow-Up

Robert returns 4 months later for an employer-sponsored flu shot. He took his medications as directed, is asymptomatic, and has had no sex partners since his office visit to you.

8) Does Robert need repeat testing for gonorrhea?

9) What are appropriate prevention counseling messages for Robert?

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